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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/578,485	06/05/2006	Peter Carmeliet	BJS-4465-10	7876

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EXAMINER	
POPA, ILEANA	

ART UNIT	PAPER NUMBER
1633	

MAIL DATE	DELIVERY MODE
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Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

Office Action Summary	Application No.	Applicant(s)	
	10/578,485	CARMELIET, PETER	
	Examiner	Art Unit	
	ILEANA POPA	1633	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

1)☐ Responsive to communication(s) filed on _____.

2a)☐ This action is **FINAL**. 2b)☒ This action is non-final.

3)☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

4)☒ Claim(s) *1-6* is/are pending in the application.

4a) Of the above claim(s) _____ is/are withdrawn from consideration.

5)☐ Claim(s) _____ is/are allowed.

6)☒ Claim(s) *1-6* is/are rejected.

7)☐ Claim(s) _____ is/are objected to.

8)☐ Claim(s) _____ are subject to restriction and/or election requirement.

Application Papers

9)☒ The specification is objected to by the Examiner.

10)☐ The drawing(s) filed on _____ is/are: a)☐ accepted or b)☐ objected to by the Examiner.

Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).

Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).

11)☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

12)☒ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).

a)☒ All b)☐ Some * c)☐ None of:

1.☐ Certified copies of the priority documents have been received.

2.☐ Certified copies of the priority documents have been received in Application No. _____.

3.☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

* See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

1) ☒ Notice of References Cited (PTO-892)
2) ☐ Notice of Draftsperson's Patent Drawing Review (PTO-948)
3) ☒ Information Disclosure Statement(s) (PTO/SB/08) Paper No(s)/Mail Date *05/23/2006; 05/05/2006*.
4) ☐ Interview Summary (PTO-413) Paper No(s)/Mail Date. _____.
5) ☐ Notice of Informal Patent Application
6) ☐ Other: _____.

DETAILED ACTION

1. Claims 1-6 are pending and under examination.

Specification

2. The disclosure is objected to because of the following informalities: this application contains sequence disclosures (p. 15, lines 22, 23, 25, and 26, p. 16, lines 22 and 23, p. 17, line 7) that are encompassed by the definitions for nucleotide sequences set forth in 37 CFR 1.821 (a)(1) and (d). However, the specification fails to comply with the requirements of 37 CFR 1.821 (a)(1) and (d), because the sequence identifiers, preceded by SEQ ID NO are missing.

Appropriate correction is required.

Claim Rejections - 35 USC § 112, second paragraph

3. The following is a quotation of the second paragraph of 35 U.S.C. 112:

> The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

4. Claim 6 is rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

Claim 6 provides for the use of a transgenic *Xenopus*, but, since the claim does not set forth any steps involved in the method/process, it is unclear what method/process applicant is intending to encompass. A claim is indefinite where it

merely recites a use without any active, positive steps delimiting how this use is actually practiced.

Claim 6 is rejected under 35 U.S.C. 101 because the claimed recitation of a use, without setting forth any steps involved in the process, results in an improper definition of a process, i.e., results in a claim which is not a proper process claim under 35 U.S.C. 101. See for example *Ex parte Dunki*, 153 USPQ 678 (Bd.App. 1967) and *Clinical Products, Ltd.* v. *Brenner*, 255 F. Supp. 131, 149 USPQ 475 (D.D.C. 1966).

Claim Rejections - 35 USC § 102

5. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

> A person shall be entitled to a patent unless –
>
> (b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

6. Claim 1 is rejected under 35 U.S.C. 102(b) as being anticipated by Beck et al. (Mechanism of Development, 1999, 88: 221-227), as evidenced by Bartel et al. (Anat Embryol, 2000, 202: 55-65).

Beck et al. teach transgenic *Xenopus* animals comprising GFP (i.e., a reporter gene) under the control of gut specific promoters (Abstract, p. 221, column 2, p. 225, column 2). It is noted that claim 1, as written, does not require the transgene to be expressed in any particular tissue or organ; the claim only recites a *Xenopus* which comprises a transgene and there is no recitation in the claim that this transgene is the gene specifically expressed in the lymphatic vessel system; since any *Xenopus* has

lymphatic vessels (see Bartel et al., p. 59, Fig. 4c), the transgenic *Xenopus* animals of Beck et al. would necessarily comprise genes which are specifically expressed in the lymphatic vessels. For these reasons, Beck et al. anticipate the instant invention.

Claim Rejections - 35 USC § 103

7. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

> (a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.

8. Claims 1-6 are rejected under 35 U.S.C. 103(a) as being unpatentable over Beck et al., in view of both Witte et al. (Microscopy Research and Technique, 2001, 55: 122-145) and Bartel et al.

** The rejection of claim 6 is based on the interpretation that the claim is drawn to a method of screening for compounds capable of modulating lymphatic vessel development.

Beck et al. teach a transgenic *Xenopus* comprising GFP (i.e., a reporter gene) under the control of mammalian tissue specific promoters such as the pancreatic-specific PDX-1 promoter, the liver-specific transthyretin promoter, and the small intestine-specific IFABP promoter, wherein the promoters drive specific GFP expression in pancreas, liver, and small intestine, respectively (claims 1 and 2) (Abstract, p. 221, column 2, p. 223, column 2, p 224, columns 1 and 2, p. 225, column 1). Beck et al. also teach a method of obtaining the transgenic *Xenopus* animals comprising GFP, wherein

GFP expression is driven by the promoters above (claim 3) and a method of visualizing the pancreas, liver, and small intestine by observing GFP expression in these transgenic *Xenopus* animals (claim 5) (p. 225, column 2, last paragraph, p. 226, column 1, third full paragraph). Although Beck et al. teach that transgenic *Xenopus* animals comprising mammalian tissue-specific promoters driving GFP expression can be generally used to study later developmental stages, such as organogenesis (Abstract, p. 221, column 2, first full paragraph, p. 225, column 2, third full paragraph), they do not specifically teach making transgenic *Xenopus* comprising GFP under the control of promoters specific for expression within the lymphatic vessels (claims 1, 3, and 4) nor do they teach using this transgenic *Xenopus* to visualize the lymphatic vessel system (claim 5) or to screen for compounds capable of modulating lymphatic vessel development (claim 6). Witte et al. teach that the lymphatic vessel development is poorly understood and suggest the use of experimental models to elucidate the mechanism of lymphatic vessel development and to develop new therapies (claim 6); Witte et al. teach using mice and not *Xenopus* as models (Abstract, p. 124, column 1, p. 127, column 1, p. 138, column 1). It would have been obvious to one of skill in the art at the time the invention was made, to modify the transgenic *Xenopus* of Beck et al. by using promoters driving specific GFP expression in lymphatic vessels (it is noted that the prior art teaches that *Xenopus* has lymphatic vessels, see) and use the resulting transgenic animals to study the development of lymphatic vessel system and to screen for agents which can modulate lymphatic vessel development as taught by Witte et al., with a reasonable expectation of success. The motivation to use transgenic *Xenopus*

and not the transgenic mice of Witte et al. is provided by Beck et al., who teach that compared to mice, *Xenopus* offers many advantages such as large number of transgenic animals in one day and visualization of GFP activity in live embryos at stages that are not accessible to mammals (p. 225, column 2, second full paragraph). The motivation to use the transgenic *Xenopus* in a method of screening is provided by Witte et al., who teach the necessity to identify agents able to modulate lymphatic vessel growth (p. 138, column 1). It is noted that, by doing so, one of skill in the art would also practice a method of visualization of the lymphatic vessel system (claim 5). One of skill in the art would have been expected to have a reasonable expectation of success in doing such because the art teaches that transgenic *Xenopus* expressing GFP or transgenes in desired tissues/organs can be successfully made and used. With respect to the limitation of comparing the effect of the tested agent by comparing treated and untreated animals (claim 6), it is noted that such a step is inherent to any method of screening for modulating agents. With respect to the limitation of the promoters recited in claim 4, Witte et al. teach that VEGFR-3 and Prox-1 are specifically expressed in the lymphatic vessels (p.). Therefore, one of skill in the art would have known to use one of these promoters to specifically express GFP in the lymphatic vessels of *Xenopus*. Thus, the claimed invention was *prima facie* obvious at the time the invention was made.

9. No claim is allowed. No claim is free of prior art.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to ILEANA POPA whose telephone number is (571)272-5546. The examiner can normally be reached on 9:00 am-5:30 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Joseph Woitach can be reached on 571-272-0739. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

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